IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

IN RE THE APPLICATION OF)
Cauwenberghs et al) Examiner:)
SERIAL NO.:)))Group Art Unit:
FILED:) () () () () () () () () () () () () ()
FOR:	DETECTION OF VON-WILLEBRANDFACTOR (vWF) ACTIVITY)))

AMENDMENT ACCOMPANYING APPLICATION

Honorable Director of Patents and Trademarks Washington, D.C. 20231

Dear Sir:

The present application is the national filing of International Application No. PCT/EP00/06345. Before calculation of the national filing fee for the United States, it is requested that the application be amended as follows:

In the claims:

Cancel claims 1-30 without prejudice and substitute new claims 31-47 as follows:

31. A method for the discrimination between von Willebrand disease (vWD) type 1 and type 2 comprising the steps of:
a) detecting vWF activity in a test sample according to the method for detecting von-Willebrand factor (vWF) activity comprising assaying a sample in the presence of a soluble form or portion of glycoprotein Ib (a) (GPlb (a)) and ristocetin, or a functionally equivalent substance.
b) determining the amount of vWF-antigen in said test sample

- c) determining the ratio between vWF-activity and vWF-antigen for said test sample; and d) comparing the under (c) obtained ratio to the range of ratios established as normal range.
- 32. The method of claim 31, wherein said detection of von-Willebrand factor (vWF) activity is carried out by detecting the formation of a complex of vWF and GPlb (a) and/or a formed complex of vWF and GPlb (a).
- 33. he method of claim 31, wherein said GPlb (a) is bound to a solid support.
- 34. The method of claim 31, wherein said GPlb (a) is bound to said solid support by a specifically reacting anti-GPlb (a) antibody.
- 35. The method of claim 31, wherein detection of von-Willebrand factor (vWF) activity is carried out by detecting the formation of a complex of vWF and GPlb (a) and/or a formed complex of vWF and GPlb and wherein said complex is bound to a solid support.
- 36. The method of claim 31, wherein detection of von-Willebrand factor (vWF) activity is carried out by detecting the formation of a complex of vWF and GPlb (a) and/or a formed complex of vWF and GPlb and wherein said complex is bound to a solid support by a specifically reacting anti-GPlb (a) antibody, by a specifically reacting anti-vWF antibody, by a specifically reacting anti-Factor VIII antibody and/or by collagen.
- 37. The method of claim 31, wherein said detection is carried out by a specifically reacting anti-vWF antibody, by a specifically reacting anti-Factor VIII antibody, by a specifically reacting anti-GPlb (a) antibody, by collagen and/or mixtures thereof.

- 38. The method of claim 31 wherein said detection is carried out by an heterogeneous or by an homogeneous assay.
- 39. The method of claim 31, wherein said detection is carried out by an heterogenous assay selected from the group of linked immuno sorbent assay (ELISA), a radioimmunoassay (RIA), an immuno radio metric assay (IRMA), a fluorescent immunoassay (FIA), a chemiluminescent immuno assay (CLIA) or an electro chemiluminescent immuno assay (ECL).
- 40. The method of claim 31, wherein said detection is carried out by an homogenous agglutination assay.
- 41. The method of claim 31 wherein the sample is selected from the group of a diluted or undiluted blood or plasma sample.
- 42. Use of a soluble form or portion of glycoprotein Ib (a) (GPlb (a)) for the discrimination between von Willebrand disease (vWD) type 1 and type 2 carrying out the steps of:
- a) detecting vWF activity in a test sample according to the method for detecting von-Willebrand factor (vWF) activity comprising assaying a sample in the presence of a soluble form or portion of glycoprotein Ib (a) (GPlb (a)) and ristocetin, or a functionally equivalent substance. b) determining the amount of vWF-antigen in said test sample;
- 43. Use of ristocetin or a functional equivalent substance of istocetin for the discrimination between von Willebrand disease (vWD) type 1 and type 2 of for carrying a) detecting vWF activity in a test sample according to the method for detecting von-Willebrand factor (vWF) activity comprising assaying a sample in the presence of a soluble form or portion of glycoprotein Ib (a) (GPlb (a)) and ristocetin, or a functionally equivalent substance. b) determining the amount of vWF-antigen in said test sample;